

The listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

1-16. (Canceled)

17. (Currently Amended) A method for the treatment of rheumatoid arthritis ~~a disease other than cancer mediated by p38~~ which comprises administering to a patient in need thereof an effective amount of a compound of formula I or a pharmaceutically acceptable salt thereof



wherein B is phenyl, pyridinyl, pyrimidinyl, pyrazinyl, pyridazinyl, naphthyl, quinolinyl, isoquinolinyl, phthalimidinyl, furyl, thieryl, pyrrolyl, imidazolyl, pyrazolyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, benzofuryl, benzothienyl, indolyl, benzopyrazolyl, benzoxazolyl, benzisoxazolyl, benzothiazolyl or benzisothiazolyl substituted by -Y-Ar; and is optionally substituted by one or more substitutents independently selected from the group consisting of halogen, up to per-halo-substitution, and X_n,

wherein n is 0-3 and each X is independently selected from the group consisting of -CN, -CO₂R⁵, -C(O)NR⁵R^{5'}, -C(O)R⁵, -NO₂, -OR⁵, -SR⁵, -NR⁵R^{5'}, -NR⁵C(O)OR⁵, -NR⁵C(O)R^{5'}, C₁-C₁₀ alkyl, C₂-C₁₀ alkenyl, C₁-C₁₀ alkoxy, C₃-C₁₀ cycloalkyl, phenyl, pyridinyl, naphthyl, isoquinolinyl, quinolinyl, up to per halo-substituted C₁-C₁₀ alkyl, up to per halo-substituted C₂-C₁₀ alkenyl, up to per halo-substituted C₁-C₁₀ alkoxy, up to per halo-substituted C₃-C₁₀ cycloalkyl, and

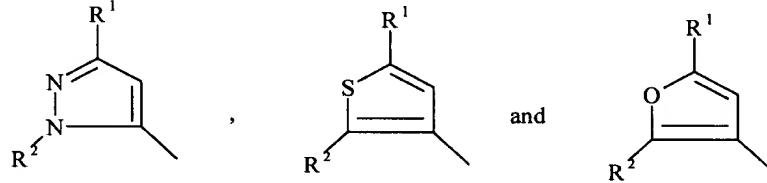
wherein R⁵ and R^{5'} are independently selected from H, C₁-C₁₀ alkyl, C₂-C₁₀ alkenyl, C₃-C₁₀ cycloalkyl, up to per-halo-substituted C₁-C₁₀ alkyl, up to per-halo-substituted C₂-C₁₀ alkenyl and up to per-halo-substituted C₃-C₁₀ cycloalkyl,

wherein Y is -O-, -S-, -N(R⁵)-, -(CH₂)_m, -C(O)-, -CH(OH)-, -(CH₂)_mO-, -NR⁵C(O)NR⁵'-, -NR⁵C(O)-, -C(O)NR⁵'-, -(CH₂)_mS-, -(CH₂)_mN(R⁵)-, -O(CH₂)_m-, -CHX^a, -CX^a₂-, -S-(CH₂)_m- and -N(R⁵)(CH₂)_m-,

m = 1-3, and X^a is halogen; and

Ar is phenyl, pyridinyl, pyrimidinyl, pyrazinyl, pyridazinyl, naphthyl, quinolinyl, isoquinolinyl, phthalimidinyl, furyl, thienyl, pyrrolyl, imidazolyl, pyrazolyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, benzofuryl, benzothienyl, indolyl, benzopyrazolyl, benzoxazolyl, benzisoxazolyl, benzothiazolyl or benzisothiazolyl, optionally substituted by halogen up to per-halo-substitution and optionally substituted by Z_{n1} , wherein $n1$ is 0 to 3 and each Z is independently selected from the group consisting of -CN, =O, -CO₂R⁵, -C(O)NR⁵R^{5'}, -C(O)-NR⁵, -NO₂, -OR⁵, -SR⁵, -NR⁵R^{5'}, -NR⁵C(O)OR^{5'}, -C(O)R⁵, -NR⁵C(O)R^{5'}, -SO₂R⁵, SO₂NR⁵R^{5'}, C₁-C₁₀ alkyl, C₁-C₁₀ alkoxy, C₃-C₁₀ cycloalkyl, up to per halo-substituted C₁-C₁₀ alkyl, and up to per halo-substituted C₃-C₁₀ cycloalkyl, and

wherein A is a heteroaryl selected from the group consisting of



wherein R¹ is selected from the group consisting of C₃-C₁₀ alkyl, C₃-C₁₀ cycloalkyl, up to per-halo-substituted C₁-C₁₀ alkyl and up to per-halo-substituted C₃-C₁₀ cycloalkyl,

wherein R² is C₆-C₁₄ aryl, C₃-C₁₄ heteroaryl, or substituted C₆-C₁₄ aryl or substituted C₃-C₁₄ heteroaryl,

wherein if R² is a substituted group, it is substituted by one or more substituents independently selected from the group consisting of halogen, up to per-halo-substitution, and V_n,

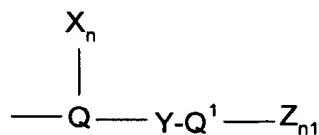
wherein n = 0-3 and each V is independently selected from the group consisting of -CN, -CO₂R⁵, -C(O)NR⁵R^{5'}, -OR⁵, -SR⁵, -NR⁵R^{5'}, -C(O)R⁵, -OC(O)NR⁵R^{5'}, -NR⁵C(O)OR^{5'}, -SO₂R⁵, -SOR⁵, -NR⁵C(O)R^{5'}, -NO₂, C₁-C₁₀ alkyl, C₃-C₁₀ cycloalkyl, C₆-C₁₄ aryl, C₃-C₁₃ heteroaryl, C₇-C₂₄ alkaryl, C₄-C₂₄ alkheteroaryl, substituted C₁-C₁₀ alkyl, substituted C₃-C₁₀ cycloalkyl, substituted C₆-C₁₄ aryl, substituted C₃-C₁₃ heteroaryl, substituted C₇-C₂₄ alkaryl and substituted C₄-C₂₄ alkheteroaryl,

where V is a substituted group, it is substituted by one or more substituents independently selected from the group consisting of halogen, up to per-halo-substitution, -CN, -CO₂R⁵, -C(O)R⁵, -C(O)NR⁵R^{5'}, -NR⁵R^{5'}, -OR⁵, -SR⁵, -NR⁵C(O)R^{5'}, -NR⁵C(O)OR^{5'} and -NO₂,

wherein R⁵ and R⁶ are each independently as defined above.

18. (Currently Amended) A method as in claim 17 wherein R² is phenyl [;] or substituted phenyl, ~~pyridinyl or substituted pyridinyl~~.

19. (Currently Amended) A method of claim 17, wherein B is



wherein

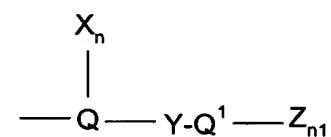
Y is as defined in claim 17,

Q and Q¹ are independently selected from the group consisting of phenyl, pyridinyl, naphthyl, pyrimidinyl, quinoline, isoquinoline, imidazole and benzothiazolyl, optionally substituted by halogen, up to per-halo substitution, and

Z and X are independently selected from the group consisting of -R⁶, -OR⁶ and -NHR⁷, wherein R⁶ is hydrogen, C₁-C₁₀-alkyl or C₃-C₁₀-cycloalkyl and R⁷ is selected from the group consisting of hydrogen, C₃-C₁₀-alkyl, and C₃-C₆-cycloalkyl wherein R⁶ and R⁷ can be substituted by halogen or up to per-halo-substitution, and

n and n1 are, each independently 0-3.

20. (Currently Amended) A method as in claim 17 19, wherein B is



wherein

Q is phenyl,

Q¹ is phenyl or pyridinyl,

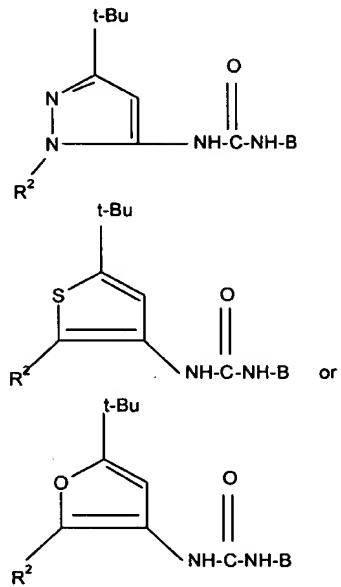
Y is -O-, -S- or -CH₂, and

X and Z are independently Cl, F, CF₃, NO₂ or CN,

n and n1 are, each independently 0-3, and

wherein Q and Q¹ are optionally substituted by one or more Cl and/or F.

21. (Previously Presented) A method as in claim 17, which comprises administering a compound of one of the formulae or a pharmaceutically acceptable salt thereof:



wherein B and R² are as defined in claim 17.

22. (Currently Amended) A method as in claim 21, wherein R² is phenyl; pyridinyl, or substituted phenyl or substituted pyridinyl.

23. (Previously Presented) A method as in claim 17, comprising administering an amount of compound of formula I effective to inhibit p38.

24. (Previously Presented) A method as in claim 17, wherein the compound of formula I displays p38 activity (IC₅₀) better than 10μM as determined by an in-vitro kinase assay.

25. (Canceled)

26. (Currently Amended) A method according to claim 30 +7, wherein R¹ is t-butyl.

27-29. (Canceled)

30. (Currently Amended) A method ~~according to claim 17, wherein the disease is for the treatment of Crohn's disease, rheumatoid arthritis, osteoarthritis, osteroporosis, asthma, septic shock, inflammatory bowel disease, or the result of host-versus-graft reactions which comprises administering to a patient in need thereof an effective amount of a compound of formula I or a pharmaceutically acceptable salt thereof~~



wherein B is phenyl, pyridinyl, pyrimidinyl, pyrazinyl, pyridazinyl, naphthyl, quinolinyl, isoquinolinyl, phthalimidinyl, furyl, thieryl, pyrrolyl, imidazolyl, pyrazolyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, benzofuryl, benzothienyl, indolyl, benzopyrazolyl, benzoxazolyl, benzisoxazolyl, benzothiazolyl or benzisothiazolyl substituted by -Y-Ar; and is optionally substituted by one or more substituents independently selected from the group consisting of halogen, up to per-halo-substitution, and X_n,

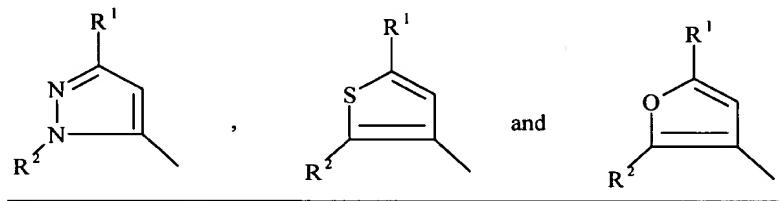
wherein n is 0-3 and each X is independently selected from the group consisting of -CN, -CO₂R⁵, -C(O)NR⁵R^{5'}, -C(O)R⁵, -NO₂, -OR⁵, -SR⁵, -NR⁵R^{5'}, -NR⁵C(O)OR⁵, -NR⁵C(O)R^{5'}, C₁-C₁₀ alkyl, C₂-C₁₀ alkenyl, C₁-C₁₀ alkoxy, C₃-C₁₀ cycloalkyl, phenyl, pyridinyl, naphthyl, isoquinolinyl, quinolinyl, up to per halo-substituted C₁-C₁₀ alkyl, up to per halo-substituted C₂-C₁₀ alkenyl, up to per halo-substituted C₁-C₁₀ alkoxy, up to per halo-substituted C₃-C₁₀ cycloalkyl, and

wherein R⁵ and R^{5'} are independently selected from H, C₁-C₁₀ alkyl, C₂-C₁₀ alkenyl, C₃-C₁₀ cycloalkyl, up to per-halo-substituted C₁-C₁₀ alkyl, up to per-halo-substituted C₂-C₁₀ alkenyl and up to per-halo-substituted C₃-C₁₀ cycloalkyl,

wherein Y is -O-, -S-, -N(R⁵)-, -(CH₂)_m, -C(O)-, -CH(OH)-, -(CH₂)_mO-, -NR⁵C(O)NR⁵'-, -NR⁵C(O)-, -C(O)NR⁵'-, -(CH₂)_mS-, -(CH₂)_mN(R⁵)-, -O(CH₂)_m-, -CHX^a, -CX^a₂-, -S-(CH₂)_m- and -N(R⁵)(CH₂)_m-,

m = 1-3, and X^a is halogen; and

Ar is phenyl, pyridinyl, pyrimidinyl, pyrazinyl, pyridazinyl, naphthyl, quinolinyl, isoquinolinyl, phthalimidinyl, furyl, thieryl, pyrrolyl, imidazolyl, pyrazolyl, oxazolyl, isoxazolyl, thiazolyl, isothiazolyl, benzofuryl, benzothienyl, indolyl, benzopyrazolyl, benzoxazolyl, benzisoxazolyl, benzothiazolyl or benzisothiazolyl, optionally substituted by halogen up to per-halo-substitution and optionally substituted by Z_{n1} , wherein $n1$ is 0 to 3 and each Z is independently selected from the group consisting of $-CN$, $=O$, $-CO_2R^5$, $-C(O)NR^5R^5'$, $-C(O)-NR^5$, $-NO_2$, $-OR^5$, $-SR^5$, $-NR^5R^5'$, $-NR^5C(O)OR^5'$, $-C(O)R^5$, $-NR^5C(O)R^5'$, $-SO_2R^5$, $SO_2NR^5R^5'$, C_1-C_{10} alkyl, C_1-C_{10} alkoxy, C_3-C_{10} cycloalkyl, up to per halo-substituted C_1-C_{10} alkyl, and up to per halo-substituted C_3-C_{10} cycloalkyl, and
wherein A is a heteroaryl selected from the group consisting of



wherein R¹ is selected from the group consisting of C_3-C_{10} alkyl, C_3-C_{10} cycloalkyl, up to per-halo-substituted C_1-C_{10} alkyl and up to per-halo-substituted C_3-C_{10} cycloalkyl,

wherein R² is C_6-C_{14} aryl, C_3-C_{14} heteroaryl, or substituted C_6-C_{14} aryl or substituted C_3-C_{14} heteroaryl,

wherein if R² is a substituted group, it is substituted by one or more substituents independently selected from the group consisting of halogen, up to per-halo-substitution, and V_n,

wherein n = 0-3 and each V is independently selected from the group consisting of -CN, $-CO_2R^5$, $-C(O)NR^5R^5'$, $-OR^5$, $-SR^5$, $-NR^5R^5'$, $-C(O)R^5$, $-OC(O)NR^5R^5'$, $-NR^5C(O)OR^5'$, $-SO_2R^5$, $-SOR^5$, $-NR^5C(O)R^5'$, $-NO_2$, C_1-C_{10} alkyl, C_3-C_{10} cycloalkyl, C_6-C_{14} aryl, C_3-C_{13} heteroaryl, C_7-C_{24} alkaryl, C_4-C_{24} alkheteroaryl, substituted C_1-C_{10} alkyl, substituted C_3-C_{10} cycloalkyl, substituted C_6-C_{14} aryl, substituted C_3-C_{13} heteroaryl, substituted C_7-C_{24} alkaryl and substituted C_4-C_{24} alkheteroaryl,

where V is a substituted group, it is substituted by one or more substituents independently selected from the group consisting of halogen, up to per-halo-substitution, -CN, $-CO_2R^5$, $-C(O)R^5$, $-C(O)NR^5R^5'$, $-NR^5R^5'$, $-OR^5$, $-SR^5$, $-NR^5C(O)R^5'$, $-NR^5C(O)OR^5'$ and $-NO_2$.

wherein R⁵ and R⁶ are each independently as defined above.

31. (Currently Amended) A method as in claim 30 4, wherein R² is phenyl.

32. (Currently Amended) A method as in claim 30 1, wherein R² is a substituted C₆-C₁₄ aryl or substituted C₃-C₁₄-heteroaryl.